

00005.001287



PATENT APPLICATION

IFW

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:)	
	:	Examiner: Not Yet Assigned
ATSUSHI OCHIAI, ET AL.)	
	:	Group Art Unit: Not Yet Assigned
Application No.: 10/568,714)	
	:	
Filed: February 17, 2006)	
	:	
For: CANCER METASTASIS)	
INHIBITOR	:	June 13, 2006

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

SUBMISSION OF MISCELLANEOUS PCT FORMS

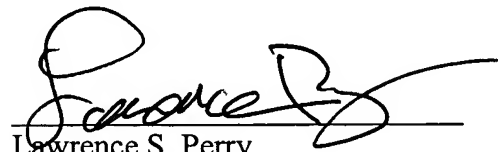
Sir:

Enclosed to complete the record and for the Examiner's convenience is Form PCT/IB/338, an English translation of the International Preliminary Report on Patentability (Form PCT/IB/373) and the Written Opinion of the International Searching Authority (Form PCT/ISA/237) in the above-identified application. The references discussed in the Written Opinion (or their patent family member equivalents) were all cited in Applicants' February 15, 2006 Information Disclosure Statement.

Entry hereof is earnestly solicited.

Applicants' undersigned attorney may be reached in our New York office by telephone at (212) 218-2100. All correspondence should continue to be directed to our below listed address.

Respectfully submitted,

A handwritten signature in black ink, appearing to read "Lawrence S. Perry", is written over a horizontal line.

Lawrence S. Perry
Attorney for Applicants
Registration. No. 31,865

FITZPATRICK, CELLA, HARPER & SCINTO
30 Rockefeller Plaza
New York, New York 10112-3801
Facsimile: (212) 218-2200

LSP\ac

NY_Main 574568_1

From the INTERNATIONAL BUREAU

PCT

NOTIFICATION OF TRANSMITTAL
OF COPIES OF TRANSLATION
OF THE INTERNATIONAL PRELIMINARY REPORT
ON PATENTABILITY
(CHAPTER I OR CHAPTER II
OF THE PATENT COOPERATION TREATY)
(PCT Rules 44bis.3(c) and 72.2)

To:

HIRAKI, Yusuke
Kamiya-cho MT Bldg. 19F
Toranomon 4-chome
Minato-ku
Tokyo, 1050001
JAPON

3-20,



Date of mailing (day/month/year) 11 May 2006 (11.05.2006)	
Applicant's or agent's file reference PH-2217-PCT	IMPORTANT NOTIFICATION
International application No. PCT/JP2004/012330	International filing date (day/month/year) 20 August 2004 (20.08.2004)
Applicant KYOWA HAKKO KOGYO CO., LTD. et al	

1. Transmittal of the translation to the applicant.



The International Bureau transmits herewith a copy of the English translation of the international preliminary report on patentability (Chapter I).



The International Bureau transmits herewith a copy of the English translation of the international preliminary report on patentability (Chapter II).

2. Transmittal of the copy of the translation to the designated or elected Offices.

The International Bureau notifies the applicant that copies of that translation have been transmitted to the following designated or elected Offices requiring such translation:

None

The following designated or elected Offices, having waived the requirement for such a transmittal at this time, will receive copies of that translation from the International Bureau only upon their request:

AE, AG, AL, AM, AP, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EA, EC, EE, EG, EP, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OA, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

3. Reminder regarding translation into (one of) the official language(s) of the elected Office(s).

The applicant is reminded that, where a translation of the international application must be furnished to an elected Office, that translation must contain a translation of any annexes to the international preliminary report on patentability (Chapter II).

It is the applicant's responsibility to prepare and furnish such translation directly to each elected Office concerned within the applicable time limit (Rule 74.1). See Volume II of the PCT Applicant's Guide for further details.

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer Yoshiko Kuwahara
Facsimile No.+41 22 740 14 35	Facsimile No.+41 22 338 90 90

PATENT COOPERATION TREATY

PCT

INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY (Chapter I of the Patent Cooperation Treaty)

(PCT Rule 44bis)

Applicant's or agent's file reference PH-2217-PCT	FOR FURTHER ACTION	See item 4 below
International application No. PCT/JP2004/012330	International filing date (<i>day/month/year</i>) 20 August 2004 (20.08.2004)	Priority date (<i>day/month/year</i>) 21 August 2003 (21.08.2003)
International Patent Classification (8th edition unless older edition indicated) See relevant information in Form PCT/ISA/237		
Applicant KYOWA HAKKO KOGYO CO., LTD.		

1. This international preliminary report on patentability (Chapter I) is issued by the International Bureau on behalf of the International Searching Authority under Rule 44 *bis*.1(a).

2. This REPORT consists of a total of 5 sheets, including this cover sheet.

In the attached sheets, any reference to the written opinion of the International Searching Authority should be read as a reference to the international preliminary report on patentability (Chapter I) instead.

3. This report contains indications relating to the following items:

- | | | |
|-------------------------------------|--------------|---|
| <input checked="" type="checkbox"/> | Box No. I | Basis of the report |
| <input type="checkbox"/> | Box No. II | Priority |
| <input type="checkbox"/> | Box No. III | Non-establishment of opinion with regard to novelty, inventive step and industrial applicability |
| <input type="checkbox"/> | Box No. IV | Lack of unity of invention |
| <input checked="" type="checkbox"/> | Box No. V | Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement |
| <input type="checkbox"/> | Box No. VI | Certain documents cited |
| <input type="checkbox"/> | Box No. VII | Certain defects in the international application |
| <input type="checkbox"/> | Box No. VIII | Certain observations on the international application |

4. The International Bureau will communicate this report to designated Offices in accordance with Rules 44bis.3(c) and 93bis.1 but not, except where the applicant makes an express request under Article 23(2), before the expiration of 30 months from the priority date (Rule 44bis .2).

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Date of issuance of this report 01 May 2006 (01.05.2006)
Facsimile No. +41 22 740 14 35	Authorized officer <div style="text-align: center; font-weight: bold; margin: 10px 0;">Yoshiko Kuwahara</div> Telephone No. +41 22 338 90 90

PATENT COOPERATION TREATY

TRANSLATION

From the
INTERNATIONAL SEARCHING AUTHORITY

PCT

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

(PCT Rule 43bis.1)

To:		<p style="font-size: 1.2em; font-weight: bold;">PCT</p> <p>WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY</p> <p>(PCT Rule 43bis.1)</p>	
		<p>Date of mailing (day/month/year) 07.12.2004</p>	
Applicant's or agent's file reference PH-2217-PCT		FOR FURTHER ACTION See paragraph 2 below	
International application No. PCT/JP2004/012330	International filing date (day/month/year) 20.08.2004	Priority date (day/month/year) 21.08.2003	
International Patent Classification (IPC) or both national classification and IPC A61K45/00, 39/395, A61P35/00			
Applicant KYOWA HAKKO KOGYO CO., LTD.			

1. This opinion contains indications relating to the following items:

- | | | |
|-------------------------------------|--------------|--|
| <input checked="" type="checkbox"/> | Box No. I | Basis of the opinion |
| <input type="checkbox"/> | Box No. II | Priority |
| <input type="checkbox"/> | Box No. III | Non-establishment of opinion with regard to novelty, inventive step and industrial applicability |
| <input type="checkbox"/> | Box No. IV | Lack of unity of invention |
| <input checked="" type="checkbox"/> | Box No. V | Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement |
| <input type="checkbox"/> | Box No. VI | Certain documents cited |
| <input type="checkbox"/> | Box No. VII | Certain defects in the international application |
| <input type="checkbox"/> | Box No. VIII | Certain observations on the international application |

2. FURTHER ACTION

If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis(b) that written opinions of this International Searching Authority will not be so considered.

If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later.

For further options, see Form PCT/ISA/220.

3. For further details, see notes to Form PCT/ISA/220.

Name and mailing address of the ISA/JP	Authorized officer
Facsimile No.	Telephone No.

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/JP2004/012330

Box No. I Basis of this opinion

1. With regard to the language, this opinion has been established on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.
☐ This opinion has been established on the basis of a translation from the original language into the following language _____, which is the language of a translation furnished for the purposes of international search (under Rule 12.3 and 23.1(b)).
2. With regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed invention, this opinion has been established on the basis of:
 - a. type of material
☒ a sequence listing
☐ table(s) related to the sequence listing
 - b. format of material
☐ in written format
☒ in computer readable form
 - c. time of filing/furnishing
☐ contained in the international application as filed.
☒ filed together with the international application in computer readable form.
☐ furnished subsequently to this Authority for the purposes of search.
3. ☒ In addition, in the case that more than one version or copy of a sequence listing and/or table(s) relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished.
4. Additional comments:

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/JP2004/012330

Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability;
citations and explanations supporting such statement

1. Statement

Novelty (N)	Claims <u>1-4</u>	YES
	Claims _____	NO
Inventive step (IS)	Claims _____	YES
	Claims <u>1-4</u>	NO
Industrial applicability (IA)	Claims <u>1-4</u>	YES
	Claims _____	NO

2. Citations and explanations:

Documents

Document 1: Cancer Research, 1998, Vol. 58, No. 15, pages
3353 to 3361

Document 2: Current Cancer Drug Targets, 2002, Vol. 2,
No. 3, pages 191 to 207

Document 3: JP 2002-512792 A (Genesense Technologies
Inc.), 8 May 2002

Explanation

Claims 1 to 4

The invention set forth in claims 1 to 4 is not disclosed in any of the documents cited in the international search report, and is therefore novel, but does not involve an inventive step in the light of documents 1 to 3 cited in the international search report.

Documents 1 and 2 indicate that metastasis is inhibited by inhibiting IGF-I receptors using methods such as administering antisense RNA or expressing dominant negatives of IGF-I receptors. Moreover, document 1 indicates that it is conceivable that both IGF-I and IGF-II which are ligands of IGF-I receptors activate cell invasion, which is an important stage, and adhesion,

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/JP2004/012330

Box No. V

Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability;
citations and explanations supporting such statement

which is the first stage of the cancer metastasis cascade, and that cancer metastasis and recurrence is prevented by targeting IGF, and that, in order for this to occur, there is a method targeting IGF-I receptors and a method targeting IGF-I and IGF-II which are ligands thereof. Document 2 indicates that an IGF-I/IGF-I receptor system promotes cancer metastasis, and that IGF-I regulates the expression of matrix metaprotease 2, which plays a central role in cancer invasion and metastasis. In addition, document 3 indicates that IGF-II contributes to the proliferation and metastasis of different types of cancer, and that IGF-II is inhibited by administering IGF-II antisense oligonucleotides, thereby suppressing cancer metastasis.

That being the case, it would be easy for a person skilled in the art to conceive of using known methods such as antibodies to inhibit the activity of both IGF-I and IGF-II which contribute to cancer metastasis, thereby promoting an improvement in the effect of suppressing cancer metastasis.